

Applicant : Kevin J. Tracey, et al.
Serial No. : 09/118,388
Filed : July 17, 1998
Page 2

insert therefor the following:

101 --Preferred compounds of formula I include, for example, 1-phenacyl-2,3-dicarboxypyridinium bromide; 1-phenacyl-2,4-dicarboxypyridinium bromide; 1-phenacyl-2,5-dicarboxypyridinium bromide; 1-phenacyl-2,6-dicarboxypyridinium bromide; 1-phenacyl-2,3-dicarboxyimidepyridinium bromide; 1-phenacyl-2,4-dicarboxyimidepyridinium bromide; 1-phenacyl-2,5-dicarboxyimidepyridinium bromide; and 1-phenacyl-2,6-dicarboxyimidepyridinium bromide--.

Delete the first full paragraph, page 35, lines 6-26 through page 36, lines 1-7, and insert therefor the following:

02 --In an alternative embodiment of the screening assay of Example 3, various concentrations of the test compound (e.g. 10-1000 μ M) are incubated with the indicator cells in presence of a fixed concentration fo 3-AP (e.g, 200 μ M). The toxicity of the test compounds may be evaluated in parallel cultures incubated without 3-AP; generally, the desired test compound will show cellular toxicity at much higher doses than thos that confer protection against 3-AP (e.g., 10-10,000-fold). The results of such tests are summarized in Table V, below.

Applicant : Kevin J. Tracey, et al.
 Serial No. : 09/118,388
 Filed : July 17, 1998
 Page 3

Table V.

Effect of test compounds on 3-AP cytotoxicity

No effect or weakly protective	Toxic or no effect	Protective (50% Effective dose; 50% Toxic dose)
Glial cell assay (HTB14)		
AP6 AP2 AP7 YA1 YA2 AP18 AP24 ascorbic acid 32P	AP9 AP12 AP19 AP20 AP23 AP28 3,5-di-tert.-butyl-4-hydroxytoluene	p27a (425 μ M; 5 mM) AP21 (100 μ M; not tested) AP22 (199 μ M; 1 mM)

wherein:

- AP6 is N-(2-phenyl-2-oxoethyl)-2-(2'-pyridine)-pyridinium bromide.
 AP2 is N-(2-phenyl-2-oxoethyl)-quinolinium bromide.
 AP7 is N-(2-phenyl-2-oxoethyl)-pyrazinium bromide.
 YA1 is 2-phenyl-2-oxoethyl-dimethylphosphonate.
 YA2 is N-(2-phenyl-2-oxoethyl)-triethylammonium bromide.
 AP18 is N-(2-phenyl-2-oxoethyl)-4-tert.-butylpyridinium bromide.
 AP24 is N-(2-phenyl-2-oxoethyl)-3-n-butylpyridinium bromide.
 34P is pyridine-3,5-dicarboxylic acid.
 AP9 is N-(2-phenyl-2-oxoethyl)-4-N,N-dimethylamino-pyridinium bromide.
 AP12 is N-(2-phenyl-2-oxoethyl)-pyrazinium bromide.
 AP19 is N-(2-phenyl-2-oxoethyl)-3-fluoropyridinium bromide.
 AP20 is N-(2-phenyl-2-oxoethyl)-4-ethylpyridinium bromide.